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FILE COVERS 1907 - 2 Nov 2006 VOL 145 ISS 19 FILE LAST UPDATED: 1 Nov 2006 (20061101/ED)

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Uploading C:\Program Files\Stnexp\Queries\fesoterodine.str

chain nodes : 23 24 25 26 27 28 8 9 10 11 12 13 14 15 16 ring nodes : 5 18 19 20 21 34 6 17 chain bonds : 10-12 11-15 11-16 12-13 12-14 5-7 7-8 7-17 8-9 9-10 10-11 19-27 22-23 23-24 24-25 24-26 27-28 28-29 ring bonds : 3-4 4-5 5-6 17-18 17-22 18-19 19-20 20-21 21-22 1-2 1-6 2-3 exact/norm bonds : 9-10 10-11 10-12 22-23 23-24 24-26 27-28 exact bonds : 8-9 11-15 11-16 12-13 12-14 19-27 24-25 28-29 5-7 7-8 7-17 normalized bonds : 4-5 5-6 17-18 17-22 18-19 19-20 20-21 21-22 1-2 1-6 2-3 3 - 4

Match level :

1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:CLASS 8:CLASS 9:CLASS 10:CLASS 11:CLASS 12:CLASS 13:CLASS 14:CLASS 15:CLASS 16:CLASS 17:Atom 18:Atom 19:Atom 20:Atom 21:Atom 22:Atom 23:CLASS 24:CLASS 25:CLASS 26:CLASS 27:CLASS 28:CLASS 29:CLASS

L1STRUCTURE UPLOADED

=> d L1

L1 HAS NO ANSWERS

L1 STR

Structure attributes must be viewed using STN Express query preparation.

=> s L1

REG1stRY INITIATED

Substance data SEARCH and crossover from CAS REGISTRY in progress... Use DISPLAY HITSTR (or FHITSTR) to directly view retrieved structures.

SAMPLE SEARCH INITIATED 12:23:18 FILE 'REGISTRY' SAMPLE SCREEN SEARCH COMPLETED -5 TO ITERATE

100.0% PROCESSED 5 ITERATIONS

0 ANSWERS

TOTAL

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **COMPLETE**

COMPLETE BATCH

PROJECTED ITERATIONS: 5 TO 234

0 TO PROJECTED ANSWERS:

0 SEA SSS SAM L1 L2

L3 0 L2

=> file caplus SINCE FILE COST IN U.S. DOLLARS

ENTRY SESSION 1.57

0.46 FULL ESTIMATED COST

SBIB ----- BIB, no citations SIBIB ----- IBIB, no citations

HIT ----- Fields containing hit terms

HITIND ----- IC, ICA, ICI, NCL, CC and index field (ST and IT)

containing hit terms

HITRN ----- HIT RN and its text modification

HITSTR ----- HIT RN, its text modification, its CA index name, and

its structure diagram

HITSEQ ----- HIT RN, its text modification, its CA index name, its

structure diagram, plus NTE and SEQ fields

FHITSTR ---- First HIT RN, its text modification, its CA index name, and

its structure diagram

FHITSEQ ---- First HIT RN, its text modification, its CA index name, its

structure diagram, plus NTE and SEQ fields

KWIC ----- Hit term plus 20 words on either side

OCC ----- Number of occurrence of hit term and field in which it occurs

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ENTER DISPLAY FORMAT (BIB):bib abs

- L5 ANSWER 1 OF 9 CAPLUS COPYRIGHT 2006 ACS on STN
- AN 2006:1133705 CAPLUS
- TI Treatment of the overactive bladder syndrome with muscarinic receptor antagonists a matter of metabolites?
- AU Michel, Martin C.; Hegde, Sharath S.
- CS Department of Pharmacology & Pharmacotherapy, Academic Medical Center, University of Amsterdam, Meibergdreef 15, Amsterdam, 1105 AZ, Neth.
- SO Naunyn-Schmiedeberg's Archives of Pharmacology (2006), 374(2), 79-85 CODEN: NSAPCC; ISSN: 0028-1298
- PB Springer
- DT Journal
- LA English

L5

Antagonists of muscarinic acetylcholine receptors, such as darifenacin, AB oxybutynin, propiverine, solifenacin, tolterodine, and trospium, are the mainstay of the treatment of the overactive bladder syndrome. Fesoterodine is a newer drug awaiting regulatory approval. We briefly review the pharmacol. activity of their metabolites and discuss how active metabolites may contribute to their efficacy and tolerability in vivo. Except for trospium, and perhaps solifenacin, all of the above drugs form active metabolites, and their presence and activity need to be taken into consideration when elucidating relationships between pharmacokinetics and pharmacodynamics of these drugs. Moreover, the ratios between parent compds. and metabolites may differ depending on genotype of the metabolizing enzymes, concomitant medication, and/or drug formulation. Differential generation of active metabolites of darifenacin or tolterodine are unlikely to influence the overall clin. profile of these drugs in a major way because the active metabolites exhibit a similar pharmacol. profile as the parent compound In contrast, metabolites of oxybutynin and propiverine may behave quant. or even qual. differently from their parent compds. and this may have an impact on the overall clin. profile of these drugs. We conclude that more comprehensive studies of drug metabolites are required for an improved understanding of their clin. effects.

```
DN
     144:156740
     Combinations of statins with bronchodilators for treatment of respiratory
TI
     disorders
     Lindmark, Bertil; Thoren, Anders Ingemar
IN
     AstraZeneca AB, Swed.; AstraZeneca UK Limited
PA
     PCT Int. Appl., 18 pp.
SO
     CODEN: PIXXD2
DT
     Patent
     English
LΑ
FAN.CNT 1
                                          APPLICATION NO.
                                                                  DATE
                         KIND
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                                         WO 2005-GB2413
     WO 2006008437
                         A1
                              20060126
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             CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,
             GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KP, KR, KZ,
             LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA,
             NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK,
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             ZA, ZM, ZW
         RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE,
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             CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM,
             KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG,
             KZ, MD, RU, TJ, TM
                                20040715
PRAI GB 2004-15789
                         Α
     The invention provides medicaments comprising combinations of
     bronchodilators, glucocorticosteroids and HMG-CoA reductase inhibitors in
     the treatment of respiratory disorders such as chronic obstructive
     pulmonary disease (COPD). For example, a metered dose inhaler contained
     per dose formoterol fumarate dihydrate 4.5 \mu g, budesonide 160 \mu g,
     rosuvastatin 1 mg, and HFA 227 50 \mu L. Also, an inhalation/oral
     combination comprised an aerosol formulation containing per dose formoterol
     fumarate dihydrate 4.5 \mu g and budesonide 160 \mu g, and a tablet
     formulation containing rosuvastatin 10 mg.
              THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT 3
              ALL CITATIONS AVAILABLE IN THE RE FORMAT
     ANSWER 3 OF 9 CAPLUS COPYRIGHT 2006 ACS on STN
L5
     2004:878361 CAPLUS
ΑN
     141:370546
DN
     Highly pure bases of 3,3-diphenyl propylamine monoesters for use in
TI
     transdermal delivery systems
     Breitenbach, Armin; Meese, Claus; Wolff, Hans-Michael; Drews, Roland
IN
     Schwarz Pharma Ag, Germany
PA
     PCT Int. Appl., 72 pp.
SO
     CODEN: PIXXD2
DT
     Patent
LΑ
     German
FAN.CNT 1
                         KIND
                                DATE
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     PATENT NO.
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                                           WO 2004-EP3567
                                20041021
                                                                   20040403
     WO 2004089872
                         A1
PΙ
         W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,
             CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,
             GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC,
             LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI,
             NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY,
             TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
         RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ,
             BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE,
             ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN,
             TD, TG
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2006:76147 CAPLUS

ΑN

	DE	1031	5917			A1	20	04111	B DE	2003	-1031	5917		2	0030	408			
	AU 2004228163				A1	20	04102	L AU	AU 2004-228163					20040403					
	CA 2505848					AA	20	04102	L CA	CA 2004-2505848					20040403				
	BR 2004006221					Α	20	05080	e BR	BR 2004-6221					20040403				
	EΡ	EP 1613584			A1	20	06011	L EP	EP 2004-725610					20040403					
		R:	ΑT,	BE,	CH,	DE,	DK, E	S, FR	GB, G	R, IT	, LI,	LU,	NL,	SE,	MC,	PT,			
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	CN	1802	345			A	20	06071	2 CN	2004	8000	9224		2	0040	403			
	JР	2006	5227	58		Т2	20	06100	JP	2006	-5049	89		2	0040	403			
	US	2006	0148	32		A1	20	06011	e us	2005	-5328	36		2	0050	426			
	NO	2005	0050	78		Α	20	05103	L NO	2005	-5078			2	0051	031			
PRAI	DE	2003	-103	1591	7	Α	20	03040	3										
	WO	2004	-EP3	567		W	20	04040	3										
OS GT	MAF	PAT	141:	3705	46														

The invention relates to a compound of general formula (I) wherein A AB represents deuterium or hydrogen, R represents a group selected from C1-6 alkyl, C3-10 cycloalkyl or Ph, which can be substituted by C1-3 alkoxy, fluorine, chlorine, bromine, iodine, nitro, amino, hydroxyl, oxo, mercapto or deuterium. The C atom marked with a * (star) can be present in an (R) configuration, in an (S)-configuration or a mixture thereof. The invention is characterized in that the above-mentioned compds. are free bases with a degree of purity of more than 97 wt %. The invention also relates to a method for the production of highly pure compds. of general formula (I) and to the use thereof in the production of medicaments. Thus (R)-2-[3-(Diisopropylamino) -1-phenylpropyl] -4-(hydroxymethyl)phenol was reacted with isobutyric acid chloride to form fesoterodine. Fesoterodine was purified via the formation of its fumaric acid salt. 1.5 G of the highly pure fesoterodine was mixed with 8.5 g silicone adhesive Bio-PSA 7-4300 and applied to a foil in order to prepare a transdermal delivery system.

RE.CNT 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

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ANSWER 4 OF 9 CAPLUS COPYRIGHT 2006 ACS on STN
L5
                 CAPLUS
ΑN
     2004:875349
DN
     142:303234
     Mucosal adjuvants and delivery systems for oral and nasal vaccination
ΤI
     Baudner, Barbara C.; Verhoel, J. Coos; Junginger, Hans E.; del Giudice,
ΑU
     Giuseppe
    IRIS Research Center, Siena, 53100, Italy
CS
     Drugs of the Future (2004), 29(7), 721-732
SO
     CODEN: DRFUD4; ISSN: 0377-8282
```

PB Prous Science

DT Journal; General Review

LA English

A review. The pillars of pharmacotherapy for overactive bladder (OAB) are AB antimuscarinic agents which inhibit bladder smooth muscle contractions through interference with acetylcholine action on muscarinic receptors of the detrusor smooth muscle. Despite the availability of different antimuscarinic compds., physicians and patients remain dissatisfied with current treatments due to adverse events and/or insufficient efficacy. Therefore, new agents with improved safety and efficacy profiles are needed for a more effective treatment of overactive bladder. Fesoterodine is a novel bladder-selective muscarinic antagonist that has shown potent antimuscarinic activity in vitro and in vivo. In multiple investigations, the agent has been shown to be safe and well tolerated in subjects of different ethnic origin, age and gender; in poor and extensive CYP2D6 metabolizers; in subjects taking concomitant medication inhibiting CYP3A4; in fed or fasted states; and in those suffering from hepatic impairment. No clin. relevant changes in heart rate, blood pressure, ECG parameters or laboratory analyses have been seen with therapeutic doses of fesoterodine in these studies. Furthermore, in a phase II clin. trial in patients with OAB, fesoterodine demonstrated rapid and significant efficacy on a variety of endpoints. The results of this trial encouraged the manufacturer (SCHWARZ PHARMA) to initiate a phase III clin. trial program for fesoterodine.

RE.CNT 169 THERE ARE 169 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

- L5 ANSWER 5 OF 9 CAPLUS COPYRIGHT 2006 ACS on STN
- AN 2004:875348 CAPLUS
- DN 142:147630
- TI Fesoterodine, an advanced antimuscarinic for the treatment of overactive bladder: a safety update
- AU Cole, Patrick
- CS Medical Information Dept., Prous Science, Barcelona, 08080, Spain
- SO Drugs of the Future (2004), 29(7), 715-720 CODEN: DRFUD4; ISSN: 0377-8282
- PB Prous Science
- DT Journal; General Review
- LA English
- The pillars of pharmacotherapy for overactive bladder (OAB) are AB A review. antimuscarinic agents which inhibit bladder smooth muscle contractions through interference with acetylcholine action on muscarinic receptors of the detrusor smooth muscle. Despite the availability of different antimuscarinic compds., physicians and patients remain dissatisfied with current treatments due to adverse events and/or insufficient efficacy. Therefore, new agents with improved safety and efficacy profiles are needed for a more effective treatment of overactive bladder. Fesoterodine is a novel bladder-selective muscarinic antagonist that has shown potent antimuscarinic activity in vitro and in vivo. multiple investigations, the agent has been shown to be safe and well tolerated in subjects of different ethnic origin, age and gender; in poor and extensive CYP2D6 metabolizers; in subjects taking concomitant medication inhibiting CYP3A4; in fed or fasted states; and in those suffering from hepatic impairment. No clin. relevant changes in heart rate, blood pressure, ECG parameters or laboratory analyses have been seen with therapeutic doses of fesoterodine in these studies. Furthermore, in a phase II clin. trial in patients with OAB, fesoterodine demonstrated rapid and significant efficacy on a variety of endpoints. The results of this trial encouraged the manufacturer (SCHWARZ PHARMA) to initiate a phase III clin. trial program for fesoterodine.
- RE.CNT 24 THERE ARE 24 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L5 ANSWER 6 OF 9 CAPLUS COPYRIGHT 2006 ACS on STN
- AN 2004:872676 CAPLUS
- DN 141:337790

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Transdermal administration of (R)-3,3-diphenylpropylamine monoesters
     Breitenbach, Armin; Meese, Claus; Wolff, Hans-Michael; Drews, Roland
IN
PA
     Schwarz Pharma Ag, Germany
SO
     PCT Int. Appl., 68 pp.
     CODEN: PIXXD2
DT
     Patent
LA
     German
FAN.CNT 1
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                                             WO 2004-EP3574
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PΙ
     WO 2004089346
                           A1
                                   20041021
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              NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY,
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              SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN,
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                                                AU 2004-228927
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     AU 2004228927
                                   20041021
                                                CA 2004-2505780
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                                                EP 2004-725614
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     EP 1530461
                            A1
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     BR 2004006212
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                                   20050816
                                                BR 2004-6212
                                                                         20040403
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20061005

20051013

20060209

20051010

20030408

20040403

CN 2004-80009176

JP 2006-504992

US 2005-533683

ZA 2005-2681

NO 2005-4644

20040403

20040403

20050401

20050426

20051010

CN 1767820

JP 2006522759

ZA 2005002681

US 2006029673

NO 2005004644

WO 2004-EP3574

MARPAT 141:337790

PRAI DE 2003-10315878

OS

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A1

Α

Α

W

The invention relates to a device for transdermally administering a compound of formula (I), wherein A represents hydrogen or deuterium, R represents a group selected among C1-6 alkyl, C3-10 cycloalkyl, or Ph, each of which can be substituted by C1-3 alkoxy, fluoride, chlorine, bromine, iodine, nitro, amino, hydroxy, oxo, mercapto, or deuterium, the C atom marked by * (asterisk) being provided in the R configuration. The invention is characterized in that the compound of general formula (I) is provided in a polymer matrix and is released at a dose of 0.5 to 20 mg per day through human skin. The invention further relates to the use of said compds. of

formula (I) for producing transdermal medicaments. Thus a silicone-based transdermal system was prepared by the hot-melt process. 8.5 G of an adhesive mixture composed of BIO-PSA 7-4300 from Dow-Corning and 5 weight/weight%

ozokerite or ceresin was heated to 150°C for 20 min until a homogeneous melt was formed. 1.5 G fesoterodine were added to the melt; the mixture was kept for addnl. 5 min at 150°C; followed by application onto a preheated foil. 5 Cm2 samples were used for dissoln. studies.

RE.CNT 6 THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

- L5 ANSWER 7 OF 9 CAPLUS COPYRIGHT 2006 ACS on STN
- AN 2004:761399 CAPLUS
- DN 141:254396
- TI Fesoterodine a new effective and well-tolerated antimuscarinic for the treatment of urgency-frequency syndrome: results of a phase 2 controlled study
- CS Chapple C1, Royal Hallamshire Hospital, UK
- SO Neurourology and Urodynamics (2004), 23(5/6), 598-599 CODEN: NEUREM; ISSN: 0733-2467
- PB Wiley-Liss, Inc.
- DT Journal
- LA English
- AB Fesoterodine as new effective and well-tolerated antimuscarinic for the treatment of urgency-frequency syndrome is studied here.
- L5 ANSWER 8 OF 9 CAPLUS COPYRIGHT 2006 ACS on STN
- AN 2003:993805 CAPLUS
- DN 140:331551
- TI Fesoterodine: Treatment of urinary incontinence muscarinic M3 antagonist
- AU Sorbera, L. A.; Castaner, J.; Lesson, P. A.
- CS Prous Science, Barcelona, 08080, Spain
- SO Drugs of the Future (2003), 28(7), 647-651 CODEN: DRFUD4; ISSN: 0377-8282
- PB Prous Science
- DT Journal; General Review
- LA English
- Urinary incontinence and overactive bladder are extremely A review. AΒ common disorders affecting up to 12 and 20 million adults in the U.S., resp. Current pharmacotherapy includes peripherally acting compds. which modulate bladder smooth muscle contraction or centrally acting agents which modulate the neurol. control of urination. Anticholinergic agents inhibit bladder smooth muscle contraction through interference with acetylcholine action on muscarinic receptors on detrusor smooth muscle. However, the first anticholinergic agents were associated with a high rate of adverse events due to nonselectivity and targeting of several muscarinic subtypes and thus other organs. The search for novel, more bladder-selective antimuscarinic agents with better tolerability was initiated. Fesoterodine is a novel selective muscarinic M3 receptor antagonist that has shown potent antimuscarinic activity in vitro and in vivo and has been selected for further development as a treatment for urinary incontinence and overactive bladder.
- RE.CNT 14 THERE ARE 14 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L5 ANSWER 9 OF 9 CAPLUS COPYRIGHT 2006 ACS on STN
- AN 2003:950829 CAPLUS
- DN 140:13084
- TI Combination of selected opioids with other active substances for use in the therapy of urinary incontinence
- IN Christoph, Thomas
- PA Grunenthal G.m.b.H., Germany
- SO PCT Int. Appl., 126 pp.

CODEN: PIXXD2

DT Patent LA German FAN.CNT 1

111110111																		
									APPLICATION NO.									
PI	WO 2003099268			A1 20031204			1	WO 2	003-	EP55		20030527						
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			PT,	RO,	RU,	SC,	SD,	SE,	SG,	SK,	SL,	TJ,	TM,	TN,	TR,	TT,	TZ,	UA,
			UG,	US,	UΖ,	VC,	VN,	YU,	ZA,	ZM,	zw							
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			KG,	ΚZ,	MD,	RU,	ТJ,	TM,	ΑT,	BE,	ВG,	CH,	CY,	CZ,	DE,	DK,	EE,	ES,
			FI,	FR,	GB,	GR,	HU,	ΙE,	IT,	LU,	MC,	NL,	PT,	RO,	SE,	SI,	SK,	TR,
			BF,	ВJ,	CF,	CG,	CI,	CM,	GΑ,	GN,	GQ,	GW,	ML,	MR,	ΝE,	SN,	TD,	TG
	DE 10224107			A1 20031211				DE 2002-10224107						20020529				
	AU 2003240717							AU 2003-240717						20030527				
	\mathbf{EP}	EP 1507520			A1 20050223			EP 2003-730120						20030527				
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	US	US 2006168942			A1	. 20060803			US 2005-545901						20050817			
PRAI	DE 2002-10224107		Α		2002	0529												
	WO	2003	-EP5	529		W		2003	0527									
os	MARPAT 140:13084																	

AB The invention discloses the use of a combination of opioids (e.g. tramadol) with other active substances for producing a drug for the treatment of urinary urgency or urinary incontinence. The invention also relates to corresponding medicaments and to a method for treating urinary urgency or urinary incontinence.

RE.CNT 9 THERE ARE 9 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

---Logging off of STN---

Executing the logoff script...

=> LOG Y

SINCE FILE TOTAL COST IN U.S. DOLLARS ENTRY SESSION 27.99 29.56 FULL ESTIMATED COST DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) SINCE FILE TOTAL ENTRY SESSION CA SUBSCRIBER PRICE -6.75 -6.75

STN INTERNATIONAL LOGOFF AT 12:25:28 ON 02 NOV 2006

Connection closed by remote host

Connecting via Winsock to STN

Welcome to STN International! Enter x:x

LOGINID: ssptalxn1621

PASSWORD:

TERMINAL (ENTER 1, 2, 3, OR ?):2

Welcome to STN International Web Page URLs for STN Seminar Schedule - N. America NEWS 1 NEWS "Ask CAS" for self-help around the clock NEWS 3 AUG 09 INSPEC enhanced with 1898-1968 archive NEWS 4 AUG 28 ADISCTI Reloaded and Enhanced NEWS 5 AUG 30 CA(SM)/CAplus(SM) Austrian patent law changes NEWS 6 SEP 11 CA/CAplus enhanced with more pre-1907 records CA/CAplus fields enhanced with simultaneous left and right NEWS 7 SEP 21 truncation CA(SM)/CAplus(SM) display of CA Lexicon enhanced NEWS 8 SEP 25 CAS REGISTRY(SM) no longer includes Concord 3D coordinates NEWS 9 SEP 25 CAS REGISTRY(SM) updated with amino acid codes for pyrrolysine NEWS 10 SEP 25 NEWS 11 SEP 28 CEABA-VTB classification code fields reloaded with new classification scheme LOGOFF HOLD duration extended to 120 minutes OCT 19 NEWS 12 NEWS 13 OCT 19 E-mail format enhanced NEWS 14 OCT 23 Option to turn off MARPAT highlighting enhancements available CAS Registry Number crossover limit increased to 300,000 in NEWS 15 OCT 23 multiple databases The Derwent World Patents Index suite of databases on STN NEWS 16 OCT 23 has been enhanced and reloaded CHEMLIST enhanced with new search and display field OCT 30 NEWS 17 NEWS EXPRESS JUNE 30 CURRENT WINDOWS VERSION IS V8.01b, CURRENT MACINTOSH VERSION IS V6.0c(ENG) AND V6.0Jc(JP), AND CURRENT DISCOVER FILE IS DATED 26 JUNE 2006. STN Operating Hours Plus Help Desk Availability NEWS HOURS Welcome Banner and News Items NEWS LOGIN For general information regarding STN implementation of IPC 8 NEWS IPC8 NEWS X25 X.25 communication option no longer available Enter NEWS followed by the item number or name to see news on that

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* * * * * * * * * * STN Columbus * *

FILE 'HOME' ENTERED AT 12:32:57 ON 02 NOV 2006

=> file reg COST IN U.S. DOLLARS

SINCE FILE TOTAL SESSION ENTRY 0.21 0.21

FULL ESTIMATED COST

FILE 'REGISTRY' ENTERED AT 12:33:15 ON 02 NOV 2006 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2006 American Chemical Society (ACS)

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http://www.cas.org/ONLINE/UG/regprops.html

=>

Uploading C:\Program Files\Stnexp\Queries\fesoterodine generic.str

chain nodes :

7 8 9 10 11 12 13 14 15 16 23 24 25 26 27 28 29 30 31

ring nodes :

1 2 3 4 5 6 17 18 19 20 21 22

chain bonds :

5-7 7-8 7-17 8-9 9-10 10-11 10-12 11-15 11-16 12-13 12-14 19-26 22-23

23-24 24-25 24-29 26-27 26-30 26-31 27-28

ring bonds :

1-2 1-6 2-3 3-4 4-5 5-6 17-18 17-22 18-19 19-20 20-21 21-22

exact/norm bonds :

9-10 10-11 10-12 22-23 23-24 24-25 26-27

exact bonds :

5-7 7-8 7-17 8-9 11-15 11-16 12-13 12-14 19-26 24-29 26-30 26-31 27-28

normalized bonds :

1-2 1-6 2-3 3-4 4-5 5-6 17-18 17-22 18-19 19-20 20-21 21-22

Match level :

1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:CLASS 8:CLASS 9:CLASS 10:CLASS 11:CLASS 12:CLASS 13:CLASS 14:CLASS 15:CLASS 16:CLASS 17:Atom 18:Atom 19:Atom 20:Atom 21:Atom 22:Atom 23:CLASS 24:CLASS 25:CLASS 26:CLASS 27:CLASS 28:CLASS 29:CLASS 30:CLASS 31:CLASS

STR

=> d L1

L1 HAS NO ANSWERS

L1

Structure attributes must be viewed using STN Express query preparation.

=> s L1

SAMPLE SEARCH INITIATED 12:33:38 FILE 'REGISTRY'
SAMPLE SCREEN SEARCH COMPLETED - 5 TO ITERATE

100.0% PROCESSED

5 ITERATIONS

0 ANSWERS

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **COMPLETE**

BATCH **COMPLETE**

PROJECTED ITERATIONS: 5 TO 234

PROJECTED ANSWERS: 0 TO 0

L2 0 SEA SSS SAM L1

=> file caplus

COST IN U.S. DOLLARS SINCE FILE TOTAL

ENTRY SESSION

FULL ESTIMATED COST 0.44 0.65

FILE 'CAPLUS' ENTERED AT 12:33:44 ON 02 NOV 2006 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS.
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=> s L2

L3 0 L2

=>

---Logging off of STN---

=>

Executing the logoff script...

=> LOG Y

COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION 0.46 1.11

FULL ESTIMATED COST

STN INTERNATIONAL LOGOFF AT 12:34:30 ON 02 NOV 2006

Connecting via Winsock to STN

Welcome to STN International! Enter x:x

LOGINID: ssptalxn1621

PASSWORD:

TERMINAL (ENTER 1, 2, 3, OR ?):2

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NEWS 2 "Ask CAS" for self-help around the clock

NEWS 3 AUG 09 INSPEC enhanced with 1898-1968 archive

NEWS 4 AUG 28 ADISCTI Reloaded and Enhanced

NEWS 5 AUG 30 CA(SM)/CAplus(SM) Austrian patent law changes

NEWS 6 SEP 11 CA/CAplus enhanced with more pre-1907 records

NEWS 7 SEP 21 CA/Caplus fields enhanced with simultaneous left and right truncation

NEWS 8 SEP 25 CA(SM)/CAplus(SM) display of CA Lexicon enhanced

NEWS 9 SEP 25 CAS REGISTRY(SM) no longer includes Concord 3D coordinates

NEWS 10 SEP 25 CAS REGISTRY(SM) updated with amino acid codes for pyrrolysine

NEWS 11 SEP 28 CEABA-VTB classification code fields reloaded with new classification scheme

NEWS 12 OCT 19 LOGOFF HOLD duration extended to 120 minutes

NEWS 13 OCT 19 E-mail format enhanced

NEWS 14 OCT 23 Option to turn off MARPAT highlighting enhancements available

NEWS 15 OCT 23 CAS Registry Number crossover limit increased to 300,000 in

multiple databases

NEWS 16 OCT 23 The Derwent World Patents Index suite of databases on STN

has been enhanced and reloaded
NEWS 17 OCT 30 CHEMLIST enhanced with new search and display field

NEWS EXPRESS JUNE 30 CURRENT WINDOWS VERSION IS V8.01b, CURRENT

MACINTOSH VERSION IS V6.0c(ENG) AND V6.0Jc(JP), AND CURRENT DISCOVER FILE IS DATED 26 JUNE 2006.

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NEWS LOGIN Welcome Banner and News Items

NEWS IPC8 For general information regarding STN implementation of IPC 8

NEWS X25 X.25 communication option no longer available

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FILE 'HOME' ENTERED AT 14:38:17 ON 02 NOV 2006

=> file reg
COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION

FULL ESTIMATED COST

0.21 0.21

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Uploading C:\Program Files\Stnexp\Queries\10532836.str

chain nodes :

7 8 9 10 11 12 13 14 15 16 17 18 19 20 21 22 23 24 25 26

ring nodes :

1 2 3 4 5 6

chain bonds :

 $5-7 \quad 7-8 \quad 7-17 \quad 8-9 \quad 9-10 \quad 10-11 \quad 10-14 \quad 11-12 \quad 11-13 \quad 14-15 \quad 14-16 \quad 17-18 \quad 17-22$

18-19 19-20 19-21 22-23 22-25 22-26 23-24

ring bonds :

1-2 1-6 2-3 3-4 4-5 5-6

exact/norm bonds :

10-11 10-14 18-19 19-21 22-23

exact bonds :

5-7 7-8 7-17 8-9 9-10 11-12 11-13 14-15 14-16 17-18 17-22 19-20 22-25

22-26 23-24

normalized bonds :

1-2 1-6 2-3 3-4 4-5 5-6

Match level :

1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:CLASS 8:CLASS 9:CLASS 10:CLASS 11:CLASS 12:CLASS 13:CLASS 14:CLASS 15:CLASS 16:CLASS 17:CLASS 18:CLASS 19:CLASS 20:CLASS 21:CLASS 22:CLASS 23:CLASS 24:CLASS 25:CLASS 26:CLASS

L1 STRUCTURE UPLOADED

=> d L1

L1 HAS NO ANSWERS

L1 STR

Structure attributes must be viewed using STN Express query preparation.

=> s L1 full

FULL SEARCH INITIATED 14:38:53 FILE 'REGISTRY'

FULL SCREEN SEARCH COMPLETED - 0 TO ITERAT

100.0% PROCESSED 0 ITERATIONS 0 ANSWERS

SEARCH TIME: 00.00.01

L2 0 SEA SSS FUL L1

=> file caplus

COST IN U.S. DOLLARS SINCE FILE TOTAL

ENTRY SESSION 166.94 167.15

FULL ESTIMATED COST

FILE 'CAPLUS' ENTERED AT 14:39:05 ON 02 NOV 2006 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2006 AMERICAN CHEMICAL SOCIETY (ACS)

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=> s L2

L3 0 L2

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=>

Executing the logoff script...

=> LOG Y

COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION 0.46 167.61

FULL ESTIMATED COST

STN INTERNATIONAL LOGOFF AT 14:39:22 ON 02 NOV 2006

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Welcome to STN International! Enter x:x

LOGINID: ssptalxn1621

PASSWORD:

TERMINAL (ENTER 1, 2, 3, OR ?):2

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NEWS 3 AUG 09 INSPEC enhanced with 1898-1968 archive

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NEWS 6 SEP 11 CA/Caplus enhanced with more pre-1907 records

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NEWS 9 SEP 25 CAS REGISTRY(SM) no longer includes Concord 3D coordinates

NEWS 10 SEP 25 CAS REGISTRY(SM) updated with amino acid codes for pyrrolysine

NEWS 11 SEP 28 CEABA-VTB classification code fields reloaded with new classification scheme

NEWS 12 OCT 19 LOGOFF HOLD duration extended to 120 minutes

NEWS 13 OCT 19 E-mail format enhanced

NEWS 14 OCT 23 Option to turn off MARPAT highlighting enhancements available

NEWS 15 OCT 23 CAS Registry Number crossover limit increased to 300,000 in multiple databases

NEWS 16 OCT 23 The Derwent World Patents Index suite of databases on STN has been enhanced and reloaded

NEWS 17 OCT 30 CHEMLIST enhanced with new search and display field

NEWS EXPRESS JUNE 30 CURRENT WINDOWS VERSION IS V8.01b, CURRENT MACINTOSH VERSION IS V6.0c(ENG) AND V6.0Jc(JP), AND CURRENT DISCOVER FILE IS DATED 26 JUNE 2006.

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NEWS LOGIN Welcome Banner and News Items

NEWS IPC8 For general information regarding STN implementation of IPC 8

NEWS X25 X.25 communication option no longer available

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FILE 'HOME' ENTERED AT 14:39:50 ON 02 NOV 2006

=> file reg
COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION 0.21 0.21

FULL ESTIMATED COST

FILE 'REGISTRY' ENTERED AT 14:40:01 ON 02 NOV 2006 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2006 American Chemical Society (ACS)

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http://www.cas.org/ONLINE/UG/regprops.html

=> s fesoterodine

L1 2 FESOTERODINE

=> d fcn

L1 ANSWER 1 OF 2 REGISTRY COPYRIGHT 2006 ACS on STN

CN Propanoic acid, 2-methyl-, 2-[(1R)-3-[bis(1-methylethyl)amino]-1phenylpropyl]-4-(hydroxymethyl)phenyl ester, (2E)-2-butenedioate (1:1) (salt) (9CI) (CA INDEX NAME)

OTHER NAMES:

CN Fesoterodine fumarate

CN SMP 8272

CN SPM 907

=> d L1

L1 ANSWER 1 OF 2 REGISTRY COPYRIGHT 2006 ACS on STN

RN 286930-03-8 REGISTRY

ED Entered STN: 21 Aug 2000

CN Propanoic acid, 2-methyl-, 2-[(1R)-3-[bis(1-methylethyl)amino]-1-

```
phenylpropyl]-4-(hydroxymethyl)phenyl ester, (2E)-2-butenedioate (1:1)
     (salt) (9CI) (CA INDEX NAME)
OTHER NAMES:
CN
     Fesoterodine fumarate
CN
     SMP 8272
CN
     SPM 907 ·
FS
     STEREOSEARCH
     C26 H37 N O3 . C4 H4 O4
MF
SR
     STN Files: ADISINSIGHT, BIOSIS, CA, CAPLUS, CBNB, IMSDRUGNEWS,
LC
       IMSPATENTS, IMSRESEARCH, PHAR, PROUSDDR, RTECS*; SYNTHLINE, TOXCENTER,
       USAN, USPATFULL
         (*File contains numerically searchable property data)
     CM
          1
     CRN
          286930-02-7
     CMF C26 H37 N O3
```

Absolute stereochemistry. Rotation (+).

CM 2

CRN 110-17-8 CMF C4 H4 O4

Double bond geometry as shown.

- 2 REFERENCES IN FILE CA (1907 TO DATE)
- 2 REFERENCES IN FILE CAPLUS (1907 TO DATE)

=> s fesoterodine isobutyrate

2 FESOTERODINE

1530 ISOBUTYRATE

L2 0 FESOTERODINE

0 FESOTERODINE ISOBUTYRATE (FESOTERODINE(W)ISOBUTYRATE)

=> s fesoteridine derivatives

0 FESOTERIDINE

165 DERIVATIVES

L3 0 FESOTERIDINE DERIVATIVES (FESOTERIDINE (W) DERIVATIVES)

=> file caplus

COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION 28.92 29.13

FULL ESTIMATED COST

FILE 'CAPLUS' ENTERED AT 14:41:33 ON 02 NOV 2006 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2006 AMERICAN CHEMICAL SOCIETY (ACS)

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http://www.cas.org/infopolicy.html

=> s fesoteridine

0 FESOTERIDINE

L4 0 FESOTERIDINE

=> s fesoterodine

L5 9 FESOTERODINE

=> d L5 1-9 all

- L5 ANSWER 1 OF 9 CAPLUS COPYRIGHT 2006 ACS on STN
- AN 2006:1133705 CAPLUS
- ED Entered STN: 30 Oct 2006
- TI Treatment of the overactive bladder syndrome with muscarinic receptor antagonists a matter of metabolites?
- AU Michel, Martin C.; Hegde, Sharath S.
- CS Department of Pharmacology & Pharmacotherapy, Academic Medical Center, University of Amsterdam, Meibergdreef 15, Amsterdam, 1105 AZ, Neth.
- SO Naunyn-Schmiedeberg's Archives of Pharmacology (2006), 374(2), 79-85 CODEN: NSAPCC; ISSN: 0028-1298
- PB Springer
- DT Journal
- LA English
- CC 1 (Pharmacology)
- Antagonists of muscarinic acetylcholine receptors, such as darifenacin, oxybutynin, propiverine, solifenacin, tolterodine, and trospium, are the mainstay of the treatment of the overactive bladder syndrome.

 Fesoterodine is a newer drug awaiting regulatory approval. We briefly review the pharmacol. activity of their metabolites and discuss how active metabolites may contribute to their efficacy and tolerability in vivo. Except for trospium, and perhaps solifenacin, all of the above drugs form active metabolites, and their presence and activity need to be taken into consideration when elucidating relationships between pharmacokinetics and pharmacodynamics of these drugs. Moreover, the ratios between parent compds. and metabolites may differ depending on genotype of the metabolizing enzymes, concomitant medication, and/or drug formulation. Differential generation of active metabolites of darifenacin or tolterodine are unlikely to influence the overall clin. profile of

these drugs in a major way because the active metabolites exhibit a similar pharmacol. profile as the parent compound In contrast, metabolites of oxybutynin and propiverine may behave quant. or even qual. differently from their parent compds. and this may have an impact on the overall clin. profile of these drugs. We conclude that more comprehensive studies of drug metabolites are required for an improved understanding of their clin. effects.

```
ANSWER 2 OF 9 CAPLUS COPYRIGHT 2006 ACS on STN
L5
ΑN
    2006:76147 CAPLUS
DN
    144:156740
ED
    Entered STN: 27 Jan 2006
    Combinations of statins with bronchodilators for treatment of respiratory
TI
    disorders
    Lindmark, Bertil; Thoren, Anders Ingemar
IN
    AstraZeneca AB, Swed.; AstraZeneca UK Limited
PA
SO
    PCT Int. Appl., 18 pp.
    CODEN: PIXXD2
DT.
    Patent
LA
    English
IC
    ICM A61K031-40
         A61K031-505; A61K031-58; A61K031-165; A61P011-00; A61P011-06;
         A61P011-08
    63-6 (Pharmaceuticals)
FAN.CNT 1
                                                                DATE
                                         APPLICATION NO.
    PATENT NO.
                       KIND DATE
                             -----
                                          _____
     _____
                        ----
                                        WO 2005-GB2413
                               20060126
                                                                 20050620
    WO 2006008437
                        A1
PΙ
         W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,
            CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,
            GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KP, KR, KZ,
            LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA,
            NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK,
            SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU,
             ZA, ZM, ZW
        RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE,
             IS, IT, LT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF,
             CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM,
             KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG,
             KZ, MD, RU, TJ, TM
PRAI GB 2004-15789
                         Α
                               20040715
CLASS
 PATENT NO.
                CLASS PATENT FAMILY CLASSIFICATION CODES
                       _____
 WO 2006008437
                ICM
                       A61K031-40
                       A61K031-505; A61K031-58; A61K031-165; A61P011-00;
                ICS
                       A61P011-06; A61P011-08
                       A61K0031-40 [ICM, 7]; A61K0031-505 [ICS, 7]; A61K0031-58
                IPCI
                        [ICS,7]; A61K0031-165 [ICS,7]; A61P0011-00 [ICS,7];
                       A61P0011-06 [ICS,7]; A61P0011-08 [ICS,7]
     The invention provides medicaments comprising combinations of
AB
     bronchodilators, glucocorticosteroids and HMG-CoA reductase inhibitors in
     the treatment of respiratory disorders such as chronic obstructive
     pulmonary disease (COPD). For example, a metered dose inhaler contained
     per dose formoterol fumarate dihydrate 4.5 \mu g, budesonide 160 \mu g,
     rosuvastatin 1 mg, and HFA 227 50 µL. Also, an inhalation/oral
     combination comprised an aerosol formulation containing per dose formoterol
     fumarate dihydrate 4.5 \mu g and budesonide 160 \mu g, and a tablet
     formulation containing rosuvastatin 10 mg.
     bronchodilator glucocorticosteroid statin respiratory disease; HMG CoA
ST
     reductase inhibitor bronchodilator respiratory disease
     Drug delivery systems
IT
        (aerosols, inhalants; combinations of statins with bronchodilators for
        treatment of respiratory disorders)
     Lung, disease
IT
```

```
(chronic obstructive pulmonary disease; combinations of statins with
       bronchodilators for treatment of respiratory disorders)
    Bronchodilators
IT
    Cholinergic antagonists
    Combination chemotherapy
    Respiratory system, disease
    β2-Adrenoceptor agonists
        (combinations of statins with bronchodilators for treatment of
        respiratory disorders)
    Glucocorticoids
IT
    RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (combinations of statins with bronchodilators for treatment of
        respiratory disorders)
    Drug delivery systems
IT
        (inhalants; combinations of statins with bronchodilators for treatment
        of respiratory disorders)
    Drug delivery systems
TT
        (powders, inhalants; combinations of statins with bronchodilators for
        treatment of respiratory disorders)
    Drug delivery systems
IT
        (tablets; combinations of statins with bronchodilators for treatment of
        respiratory disorders)
                                                   100-76-5D, Quinuclidine,
     50-24-8, Prednisolone
                            53-03-2, Prednisone
IT
              124-94-7, Triamcinolone 596-51-0 3385-03-3, Flunisolide
                                 25990-43-6, Mepenzolate
                                                           51333-22-3,
     4419-39-0, Beclomethasone
                                            60205-81-4, Ipratropium
                  60135-22-0, Flumoxonide
     Budesonide
                             73573-88-3, Mevastatin
                                                       75330-75-5, Lovastatin
     73573-87-2, Formoterol
     79902-63-9, Simvastatin 81093-37-0, Pravastatin 81732-65-2, Bambuterol
     85197-77-9, Tipredane 89365-50-4, Salmeterol 90566-53-3, Fluticasone
     93957-54-1, Fluvastatin 99571-64-9, Oxitropium 105102-22-5, Mometasone
                                                         126544-47-6,
     120815-74-9, Butixocort 124937-51-5, Tolterodine
                                             133099-04-4, Darifenacin
     Ciclesonide 129260-79-3, Loteprednol
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                                                                   137888-11-0,
     134523-00-5, Atorvastatin
               144459-70-1, Rofleponide
                                         145599-86-6, Cerivastatin
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     170105-16-5, Imidafenacin 182069-13-2, ETIPREDNOL
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     Formoterol fumarate dihydrate
                                     186691-13-4, Tiotropium
                                286930-02-7, Fesoterodine
     242478-37-1, Solifenacin
     287714-41-4, Rosuvastatin 397864-44-7, 6α,9α-Difluoro-
     17\alpha-[(2-furanylcarbonyl)oxy]-11\beta-hydroxy-16\alpha-methyl-3-oxo-
     androsta-1,4-diene-17β-carbothioic acid S-fluoromethyl ester
                   452339-68-3, 3-[4-[[6-[[(2R)-2-Hydroxy-2-[4-hydroxy-3-
     (hydroxymethyl)phenyl]ethyl]amino]hexyl]oxy]butyl]benzenesulfonamide
                                             867022-63-7
     463934-65-8 678160-57-1, Zoticasone
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (combinations of statins with bronchodilators for treatment of
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     9028-35-7, HMG-CoA reductase
IT
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
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IT
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        (pitavastatin; combinations of statins with bronchodilators for
        treatment of respiratory disorders)
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     141:370546
ED
     Entered STN: 22 Oct 2004
     Highly pure bases of 3,3-diphenyl propylamine monoesters for use in
TТ
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transdermal delivery systems
    Breitenbach, Armin; Meese, Claus; Wolff, Hans-Michael; Drews, Roland
IN
PA
    Schwarz Pharma Ag, Germany
    PCT Int. Appl., 72 pp.
SO
    CODEN: PIXXD2
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    ICM C07C217-62
IC
    ICS A61K031-135; C07C213-10; A61P013-00
    63-6 (Pharmaceuticals)
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                       C07C217-62
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                ICM
                       A61K031-135; C07C213-10; A61P013-00
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                        A61P0013-00 [I,A]
                        A61K009/70E; A61K031/135; C07C213/10; C07C217/62
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                        [I,A]; A61P0013-10 [I,A]; A61P0013-00 [I,C*];
                        A61K0009-70 [I,A]; A61K0047-32 [I,A]; C07B0053-00 [N,A]
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                        4C076/EE48Q; 4C076/FF31; 4C076/FF63; 4C076/FF68;
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                        4H006/BB15; 4H006/BB16; 4H006/BB17; 4H006/BB31;
                        4H006/BC16; 4H006/BE12; 4H006/BE13; 4H006/BJ50;
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                        [I,A]; A61K0031-21 [I,C*]
                        514/540.000; 560/136.000
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OS
     MARPAT 141:370546
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The invention relates to a compound of general formula (I) wherein A AB represents deuterium or hydrogen, R represents a group selected from C1-6 alkyl, C3-10 cycloalkyl or Ph, which can be substituted by C1-3 alkoxy, fluorine, chlorine, bromine, iodine, nitro, amino, hydroxyl, oxo, mercapto or deuterium. The C atom marked with a * (star) can be present in an (R) configuration, in an (S)-configuration or a mixture thereof. The invention is characterized in that the above-mentioned compds. are free bases with a degree of purity of more than 97 wt %. The invention also relates to a method for the production of highly pure compds. of general formula (I) and to the use thereof in the production of medicaments. Thus (R)-2-[3-(Diisopropylamino) -1-phenylpropyl] -4-(hydroxymethyl)phenol was reacted with isobutyric acid chloride to form fesoterodine. Fesoterodine was purified via the formation of its fumaric acid salt. 1.5 G of the highly pure fesoterodine was mixed with 8.5 g silicone adhesive Bio-PSA 7-4300 and applied to a foil in order to prepare a transdermal delivery system.

fesoterodine purifn monoester transdermal delivery system ST

IT Ion exchangers

(basic; highly pure bases of 3,3-di-Ph propylamine monoesters for use in transdermal delivery systems)

IT Bladder

> (detrusor muscle, hyperactivity of; highly pure bases of 3,3-di-Ph propylamine monoesters for use in transdermal delivery systems)

IT Adhesives

Chirality

Crystallization

Dissolution

(highly pure bases of 3,3-di-Ph propylamine monoesters for use in transdermal delivery systems)

IT Amines, reactions

Bicarbonates

RL: RCT (Reactant); RACT (Reactant or reagent)

(highly pure bases of 3,3-di-Ph propylamine monoesters for use in transdermal delivery systems)

IT Bladder, disease

(incontinence; highly pure bases of 3,3-di-Ph propylamine monoesters for use in transdermal delivery systems)

IT Urinary system, disease

(nocturia; highly pure bases of 3,3-di-Ph propylamine monoesters for use in transdermal delivery systems)

IT Bladder, disease

(pollakisuria; highly pure bases of 3,3-di-Ph propylamine monoesters for use in transdermal delivery systems)

IT Amines, reactions

> RL: RCT (Reactant); RACT (Reactant or reagent) (polyamines, nonpolymeric; highly pure bases of 3,3-di-Ph propylamine monoesters for use in transdermal delivery systems)

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IT
     Drug delivery systems
        (transdermal; highly pure bases of 3,3-di-Ph propylamine monoesters for
        use in transdermal delivery systems)
     Drug delivery systems
TT
        (transmucosal; highly pure bases of 3,3-di-Ph propylamine monoesters
        for use in transdermal delivery systems)
     60-29-7, Diethyl ether, uses
                                    75-09-2, Dichloromethane, uses
                                                                      78-93-3,
IT
     Ethylmethylketone, uses 108-88-3, Toluene, uses
                                                         141-78-6,
                         1634-04-4, tert. Butylmethyl ether
     Ethylacetate, uses
     RL: NUU (Other use, unclassified); USES (Uses)
        (highly pure bases of 3,3-di-Ph propylamine monoesters for use in
        transdermal delivery systems)
IT
     286930-02-7P, Fesoterodine
     RL: PRP (Properties); RCT (Reactant); SPN (Synthetic preparation); THU
     (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT
     (Reactant or reagent); USES (Uses)
        (highly pure bases of 3,3-di-Ph propylamine monoesters for use in
        transdermal delivery systems)
     504415-91-2P, Bio-PSA 7-4300
IT
     RL: PRP (Properties); SPN (Synthetic preparation); PREP (Preparation)
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IT
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     BIOL (Biological study); PREP (Preparation); USES (Uses)
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                                         110-17-8, Fumaric acid, reactions
     79-30-1, Isobutyric acid chloride
IT
     207679-81-0
     RL: RCT (Reactant); RACT (Reactant or reagent)
        (highly pure bases of 3,3-di-Ph propylamine monoesters for use in
        transdermal delivery systems)
     5586-73-2D, 3,3-Diphenyl propylamine, monoesters
IT
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (highly pure bases of 3,3-di-Ph propylamine monoesters for use in
        transdermal delivery systems)
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     2004:875349 CAPLUS
ΑN
     142:303234
DN
     Entered STN: 22 Oct 2004
ED
     Mucosal adjuvants and delivery systems for oral and nasal vaccination
ΤI
     Baudner, Barbara C.; Verhoel, J. Coos; Junginger, Hans E.; del Giudice,
ΑU
     Giuseppe
     IRIS Research Center, Siena, 53100, Italy
CS
     Drugs of the Future (2004), 29(7), 721-732
SO
     CODEN: DRFUD4; ISSN: 0377-8282
     Prous Science
PB
DT
     Journal; General Review
LΑ
     English
     63-0 (Pharmaceuticals)
CC
     Section cross-reference(s): 15
     A review. The pillars of pharmacotherapy for overactive bladder (OAB) are
AB
     antimuscarinic agents which inhibit bladder smooth muscle contractions
     through interference with acetylcholine action on muscarinic receptors of
     the detrusor smooth muscle. Despite the availability of different
     antimuscarinic compds., physicians and patients remain dissatisfied with
     current treatments due to adverse events and/or insufficient efficacy.
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Therefore, new agents with improved safety and efficacy profiles are

needed for a more effective treatment of overactive bladder. Fesoterodine is a novel bladder-selective muscarinic antagonist that has shown potent antimuscarinic activity in vitro and in vivo. In multiple investigations, the agent has been shown to be safe and well tolerated in subjects of different ethnic origin, age and gender; in poor and extensive CYP2D6 metabolizers; in subjects taking concomitant medication inhibiting CYP3A4; in fed or fasted states; and in those suffering from hepatic impairment. No clin. relevant changes in heart rate, blood pressure, ECG parameters or laboratory analyses have been seen with therapeutic doses of fesoterodine in these studies. Furthermore, in a phase II clin. trial in patients with OAB, fesoterodine demonstrated rapid and significant efficacy on a variety of endpoints. The results of this trial encouraged the manufacturer (SCHWARZ PHARMA) to initiate a phase III clin. trial program for fesoterodine.

- ST review mucosa adjuvant oral nasal vaccine
- IT Immunostimulants

(adjuvants; mucosal adjuvants and delivery systems for oral and nasal vaccination)

IT Muscarinic antagonists

Vaccines

(mucosal adjuvants and delivery systems for oral and nasal vaccination)

IT Drug delivery systems

(nasal; mucosal adjuvants and delivery systems for oral and nasal vaccination)

IT Drug delivery systems

(oral; mucosal adjuvants and delivery systems for oral and nasal vaccination)

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     2004:875348 CAPLUS
AN
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ED
     Entered STN: 22 Oct 2004
TI
     Fesoterodine, an advanced antimuscarinic for the treatment of
     overactive bladder: a safety update
AU
     Cole, Patrick
     Medical Information Dept., Prous Science, Barcelona, 08080, Spain
CS
SO
     Drugs of the Future (2004), 29(7), 715-720
     CODEN: DRFUD4; ISSN: 0377-8282
PB
     Prous Science
     Journal; General Review
DT
LΆ
     English
     1-0 (Pharmacology)
CC
     A review. The pillars of pharmacotherapy for overactive bladder (OAB) are
AΒ
     antimuscarinic agents which inhibit bladder smooth muscle contractions
     through interference with acetylcholine action on muscarinic receptors of
     the detrusor smooth muscle. Despite the availability of different
     antimuscarinic compds., physicians and patients remain dissatisfied with
     current treatments due to adverse events and/or insufficient efficacy.
     Therefore, new agents with improved safety and efficacy profiles are
     needed for a more effective treatment of overactive bladder.
     Fesoterodine is a novel bladder-selective muscarinic antagonist
     that has shown potent antimuscarinic activity in vitro and in vivo.
     multiple investigations, the agent has been shown to be safe and well
     tolerated in subjects of different ethnic origin, age and gender; in poor
     and extensive CYP2D6 metabolizers; in subjects taking concomitant
     medication inhibiting CYP3A4; in fed or fasted states; and in those
     suffering from hepatic impairment. No clin. relevant changes in heart
     rate, blood pressure, ECG parameters or laboratory analyses have been seen with
     therapeutic doses of fesoterodine in these studies.
     Furthermore, in a phase II clin. trial in patients with OAB,
     fesoterodine demonstrated rapid and significant efficacy on a
     variety of endpoints. The results of this trial encouraged the
     manufacturer (SCHWARZ PHARMA) to initiate a phase III clin. trial program
     for fesoterodine.
     review fesoterodine antimuscarinic overactive bladder
ST
IT
     Combination chemotherapy
     Drug interactions
     Human
     Muscarinic antagonists
        (advanced antimuscarinic fesoterodine for treatment of
        overactive bladder)
IT
     Bladder, disease
        (hyperreflexia; advanced antimuscarinic fesoterodine for
        treatment of overactive bladder)
     286930-02-7, Fesoterodine
IT
     RL: ADV (Adverse effect, including toxicity); PAC (Pharmacological
     activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (advanced antimuscarinic fesoterodine for treatment of
        overactive bladder)
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RE
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     2004:872676 CAPLUS
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     141:337790
ED
     Entered STN: 21 Oct 2004
     Transdermal administration of (R)-3,3-diphenylpropylamine monoesters
ΤI
     Breitenbach, Armin; Meese, Claus; Wolff, Hans-Michael; Drews, Roland
     Schwarz Pharma Ag, Germany
PA
SO
     PCT Int. Appl., 68 pp.
     CODEN: PIXXD2
DT
     Patent
     German
LΑ
     ICM A61K009-70
IC
     ICS A61K031-403; C07C219-26
CC
     63-6 (Pharmaceuticals)
     Section cross-reference(s): 1
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CLASS
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                        A61K0009-70 [ICM,7]; A61K0031-403 [ICS,7]; C07C0219-26
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                        4C076/FF68; 4C206/AA01; 4C206/AA02; 4C206/DB03;
                        4C206/DB04; 4C206/DB57; 4C206/MA01; 4C206/MA04;
                        4C206/MA52; 4C206/MA83; 4C206/NA11; 4C206/NA12;
                        4C206/ZA81
                        A61K [ICS,7]; C07C [ICS,7] .
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     MARPAT 141:337790
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GI

The invention relates to a device for transdermally administering a compound AB of formula (I), wherein A represents hydrogen or deuterium, R represents a group selected among C1-6 alkyl, C3-10 cycloalkyl, or Ph, each of which can be substituted by C1-3 alkoxy, fluoride, chlorine, bromine, iodine, nitro, amino, hydroxy, oxo, mercapto, or deuterium, the C atom marked by * (asterisk) being provided in the R configuration. The invention is characterized in that the compound of general formula (I) is provided in a polymer matrix and is released at a dose of 0.5 to 20 mg per day through human skin. The invention further relates to the use of said compds. of formula (I) for producing transdermal medicaments. Thus a silicone-based transdermal system was prepared by the hot-melt process. 8.5 G of an adhesive mixture composed of BIO-PSA 7-4300 from Dow-Corning and 5

weight/weight%

ozokerite or ceresin was heated to 150°C for 20 min until a homogeneous melt was formed. 1.5 G fesoterodine were added to the melt; the mixture was kept for addnl. 5 min at 150°C; followed by application onto a preheated foil. 5 Cm2 samples were used for dissoln. studies.

transdermal diphenylpropylamine monoester Fesoterodineincontinence ST

IT Isoprene-styrene rubber

RL: PEP (Physical, engineering or chemical process); PYP (Physical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)

(block, triblock; transdermal administration of (R)-3,3diphenylpropylamine monoesters)

IT Bladder, disease

(incontinence; transdermal administration of (R)-3,3-

diphenylpropylamine monoesters)

IT Urinary system, disease

(nocturia; transdermal administration of (R)-3,3-diphenylpropylamine monoesters)

IT Paraffin oils

RL: PEP (Physical, engineering or chemical process); PYP (Physical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)

(ondina oil; transdermal administration of (R)-3,3-diphenylpropylamine monoesters)

TT Dissolution

Human

Hydrophilicity

Ozocerite

(transdermal administration of (R)-3,3-diphenylpropylamine monoesters)

IT Ceresin

TT

RL: PEP (Physical, engineering or chemical process); PYP (Physical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)

(transdermal administration of (R)-3,3-diphenylpropylamine monoesters) Polyoxyalkylenes, biological studies

```
RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (transdermal administration of (R)-3,3-diphenylpropylamine monoesters)
IT
     Drug delivery systems
        (transdermal; transdermal administration of (R)-3,3-diphenylpropylamine
        monoesters)
     Urinary system, disease
IT
        (urinary frequency; transdermal administration of (R)-3,3-
        diphenylpropylamine monoesters)
TI
                 700836-36-8D, block, triblock
     RL: PEP (Physical, engineering or chemical process); PYP (Physical
     process); THU (Therapeutic use); BIOL (Biological study); PROC (Process);
     USES (Uses)
        (isoprene-styrene rubber; transdermal administration of
        (R)-3,3-diphenylpropylamine monoesters)
IT
     286930-02-7P, Fesoterodine
     RL: PEP (Physical, engineering or chemical process); PRP (Properties); PYP
     (Physical process); SPN (Synthetic preparation); THU (Therapeutic use);
     BIOL (Biological study); PREP (Preparation); PROC (Process); USES (Uses)
        (transdermal administration of (R)-3,3-diphenylpropylamine monoesters)
     1617-18-1, Ethylvinylacetate 198292-68-1, DuroTak 387-2287
IT
     346577-82-0, Regalite R 1090
                                  504415-91-2, BIO-PSA 7-4300
     RL: PEP (Physical, engineering or chemical process); PYP (Physical
     process); THU (Therapeutic use); BIOL (Biological study); PROC (Process);
     USES (Uses)
        (transdermal administration of (R)-3,3-diphenylpropylamine monoesters)
IT
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              THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT
RE
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     ANSWER 7 OF 9 CAPLUS COPYRIGHT 2006 ACS on STN
L5
     2004:761399 CAPLUS
AN
DN
     141:254396
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ED
     Fesoterodine a new effective and well-tolerated antimuscarinic
TI
     for the treatment of urgency-frequency syndrome: results of a phase 2
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     Chapple C1, Royal Hallamshire Hospital, UK
CS
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PΒ
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LΑ
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     1-11 (Pharmacology)
     Fesoterodine as new effective and well-tolerated antimuscarinic
AB
     for the treatment of urgency-frequency syndrome is studied here.
     antimuscarinic fesoterodine urgency frequency syndrome urinary
st
     incontinence
TT
     Human
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Muscarinic antagonists (antimuscarinic fesoterodine for treatment of urgency-frequency syndrome) IT Bladder, disease (incontinence; antimuscarinic fesoterodine for treatment of urgency-frequency syndrome) IT Disease, animal (urgency-frequency syndrome; antimuscarinic fesoterodine for treatment of urgency-frequency syndrome) 286930-02-7, Fesoterodine IT RL: ADV (Adverse effect, including toxicity); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (antimuscarinic fesoterodine for treatment of urgency-frequency syndrome) ANSWER 8 OF 9 CAPLUS COPYRIGHT 2006 ACS on STN L_5 AN 2003:993805 CAPLUS 140:331551 DNED Entered STN: 22 Dec 2003 Fesoterodine: Treatment of urinary incontinence muscarinic M3 TI antagonist Sorbera, L. A.; Castaner, J.; Lesson, P. A. ΑU CS Prous Science, Barcelona, 08080, Spain SO Drugs of the Future (2003), 28(7), 647-651 CODEN: DRFUD4; ISSN: 0377-8282 PB Prous Science Journal; General Review DTLA English CC 1-0 (Pharmacology) A review. Urinary incontinence and overactive bladder are extremely AB common disorders affecting up to 12 and 20 million adults in the U.S., resp. Current pharmacotherapy includes peripherally acting compds. which modulate bladder smooth muscle contraction or centrally acting agents which modulate the neurol. control of urination. Anticholinergic agents inhibit bladder smooth muscle contraction through interference with acetylcholine action on muscarinic receptors on detrusor smooth muscle. However, the first anticholinergic agents were associated with a high rate of adverse events due to nonselectivity and targeting of several muscarinic subtypes and thus other organs. The search for novel, more bladder-selective antimuscarinic agents with better tolerability was initiated. Fesoterodine is a novel selective muscarinic M3 receptor antagonist that has shown potent antimuscarinic activity in vitro and in vivo and has been selected for further development as a treatment for urinary incontinence and overactive bladder. review fesoterodine urine incontinence muscarinic M3 antagonist ST Muscarinic antagonists IT (M3; fesoterodine treatment of urinary incontinence as muscarinic M3 antagonist) IT Bladder, disease (incontinence; fesoterodine treatment of urinary incontinence as muscarinic M3 antagonist) 286930-02-7, Fesoterodine IT RL: DMA (Drug mechanism of action); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (fesoterodine treatment of urinary incontinence as muscarinic M3 antagonist) THERE ARE 14 CITED REFERENCES AVAILABLE FOR THIS RECORD RE.CNT 14 RE (1) Andersson, K; BJU Int 1999, V84, P923 CAPLUS (2) Andersson, K; Bailliere's Best Pract Res Clin Obstet Gynaecol 2000, V14,

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    ANSWER 9 OF 9 CAPLUS COPYRIGHT 2006 ACS on STN
L5
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    2003:950829 CAPLUS
    140:13084
DN
ED
    Entered STN: 07 Dec 2003
    Combination of selected opioids with other active substances for use in
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    the therapy of urinary incontinence
    Christoph, Thomas
IN
PA
    Grunenthal G.m.b.H., Germany
SO
     PCT Int. Appl., 126 pp.
    CODEN: PIXXD2
DT
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    German
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     ICM A61K031-135
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     ICS A61K031-137; A61K031-485
     1-12 (Pharmacology)
     Section cross-reference(s): 63
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                        060/276.000; 060/285.000
                        A61K031/135; A61K031/137; A61K031/485
                 ECLA
os
     MARPAT 140:13084
     The invention discloses the use of a combination of opioids (e.g.
AΒ
     tramadol) with other active substances for producing a drug for the
     treatment of urinary urgency or urinary incontinence. The invention also
     relates to corresponding medicaments and to a method for treating urinary
     urgency or urinary incontinence.
     incontinence urinary treatment opioid drug combination; urinary urge
ST
     treatment opioid drug combination; tramadol drug combination urinary
     incontinence urge
     Bladder, disease
IT
        (incontinence; opioid combination with other active substances for
        treatment of urinary incontinence)
     Drug delivery systems
IT
        (injections; opioid combination with other active substances for
        treatment of urinary incontinence)
     Drug delivery systems
IT
        (opioid combination with other active substances for treatment of
        urinary incontinence)
     Bladder
IT
        (urinary urge; opioid combination with other active substances for
        treatment of urinary incontinence)
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                                          21363-18-8, Viminol
                                                                 27203-92-5,
     Etorphine
                                          51931-66-9, Tilidine
                                                                 52485-79-7,
                42408-82-2, Butorphanol
     Tramadol
                                            54340-58-8, Meptazinol
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                    138853-73-3
     RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (Combination of selected opioids with other active substances for use
        in the therapy of urinary incontinence)
IT
     186033-14-7, NS 8
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     (Biological study); USES (Uses)
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     52-28-8, Codeine phosphate
IT
               93413-69-5, Venlafaxine 142155-43-9, Cizolirtine
     158836-71-6, Nitro-Flurbiprofen 174636-32-9, Talnetant
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RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(opioid combination with other active substances for treatment of urinary incontinence)

RE.CNT 9 THERE ARE 9 CITED REFERENCES AVAILABLE FOR THIS RECORD RE

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=> FIL MARPAT

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FILE CONTENT: 1961-PRESENT VOL 145 ISS 18 (20061027/ED)

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7108861 19 SEP 2006
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IT

Drug delivery systems.

urinary incontinence)

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ANSWER 1 MARPAT COPYRIGHT 2006 ACS on STN
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     Combination of selected opioids with other active substances for use in
TI
     the therapy of urinary incontinence
IN
     Christoph, Thomas
     Grunenthal G.m.b.H., Germany
SO
     PCT Int. Appl., 126 pp.
     CODEN: PIXXD2
DT
     Patent
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     German
IC
     ICM A61K031-135
     ICS A61K031-137; A61K031-485
     1-12 (Pharmacology)
CC
     Section cross-reference(s): 63
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     The invention discloses the use of a combination of opioids (e.g.
AB
     tramadol) with other active substances for producing a drug for the
     treatment of urinary urgency or urinary incontinence. The invention also
     relates to corresponding medicaments and to a method for treating urinary
     urgency or urinary incontinence.
     incontinence urinary treatment opioid drug combination; urinary urge
ST
     treatment opioid drug combination; tramadol drug combination urinary
     incontinence urge
TT
     Bladder, disease
        (incontinence; opioid combination with other active substances for
        treatment of urinary incontinence)
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     Drug delivery systems
        (injections; opioid combination with other active substances for
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(opioid combination with other active substances for treatment of

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                   433686-05-6
                                 433686-06-7
                                               433686-07-8
                                                             433936-14-2
     433686-04-5
                                               502616-18-4
                                                             502616-19-5
     433936-20-0
                   433936-23-3
                                 433936-24-4
                                 502616-23-1
                                               630046-59-2
                                                             630395-07-2, SL
                  502616-22-0
     502616-20-8
             630395-09-4, DRP 001
     RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (opioid combination with other active substances for treatment of
        urinary incontinence)
              THERE ARE 9 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT
(1) Durand, A; PRESSE MEDICALE 2000, V29(16), P917
(2) Gruenenthal Gmbh; DE 19947747 A 2001 CAPLUS
(3) Kroner, B; JOURNAL OF GERIATRIC DRUG THERAPY 1992, V7(1), P23
(4) Malinovsky, J; ANESTHESIA AND ANALGESIA 1998, V87(2), P456 CAPLUS
(5) McNutt, R; US 5658908 A 1997 CAPLUS
(6) Novosis Pharma Ag; EP 1072260 A 2001 CAPLUS
(7) Palmer, K; GASTROENTEROLOGY 1980, V79(6), P1272 MEDLINE
(8) Pandita, R; NEUROUROLOGY AND URODYNAMICS, 31st Annual Meeting of the
    International Continence Society 2001, V20(4), P439
(9) Ripple, M; AMERICAN JOURNAL OF FORENSIC MEDICINE AND PATHOLOGY 2000,
```

V21(4), P370 MEDLINE

G1 = OH / F / Cl / H / 22

- G2 = carbon chain <containing 1-3 C> (opt. substd.) / (Specifically claimed: Me)
- G3 = carbon chain <containing 1-4 C> (opt. substd.) / (Specifically claimed: Me / Et / Bu-n / Bu-t)
- G4 = 27 / cycloalkylene <containing 4-7 C> (opt. substd.) / (Specifically claimed: 99)

- G5 = H / carbon chain <containing 1-4 C> (opt. substd.) / (Specifically claimed: Me / Et / Pr-i / Bu-t)
- G6 = 2-136 3-134 4-135 6-12 / 46-136 47-134 48-135 49-12 / 55-136 59-134 58-135 60-12

G7 = H / F / Cl / Br / I / 29 / OH / SH / 33 / OCF3 / NH2 / 35 / SO2Me / SO2CF3 / CN / 91 / NO2 / CONH2 / 41 / carbon chain <containing 1-6 C> (opt. substd.) / Ph (opt. substd.)

G25 = carbon chain <containing 1-5 C> (opt. substd.)

G26 = 148 / 151

$$G1$$
 C CH_2 Me $G1$ CH_2 Me $G3$ $G4$ N $G2$ Me Me

G27 = alkyl <containing 1-4 C> / CH2Ph / CF3 / OH / OCH2Ph / alkoxy <containing 1-4 C> / Cl / F / H /

(Specifically claimed: Me)

Patent location:

claim 1

Note:

and/or physiologically acceptable salts

Stereochemistry:

and enantiomers, diastereomers and mixtures

=> SET NOTICE LOGIN DISPLAY

NOTICE SET TO OFF FOR DISPLAY COMMAND SET COMMAND COMPLETED

=> FIL STNGUIDE

COST IN U.S. DOLLARS
SINCE FILE TOTAL
ENTRY SESSION
FULL ESTIMATED COST
8.14
69.72

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)

SINCE FILE TOTAL
ENTRY SESSION

CA SUBSCRIBER PRICE

-0.71
-7.46

FILE 'STNGUIDE' ENTERED AT 14:43:44 ON 02 NOV 2006
USE IS SUBJECT TO THE TERMS OF YOUR CUSTOMER AGREEMENT
COPYRIGHT (C) 2006 AMERICAN CHEMICAL SOCIETY, JAPAN SCIENCE
AND TECHNOLOGY CORPORATION, AND FACHINFORMATIONSZENTRUM KARLSRUHE

FILE CONTAINS CURRENT INFORMATION.
LAST RELOADED: Oct 27, 2006 (20061027/UP).

= >

---Logging off of STN---

=>

Executing the logoff script...

=> LOG Y

---Logging off of STN---

COST IN U.S. DOLLARS SINCE FILE TOTAL ENTRY SESSION TULL ESTIMATED COST 0.30 70.02

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)

SINCE FILE TOTAL ENTRY SESSION

CA SUBSCRIBER PRICE

0.00

-7.46

STN INTERNATIONAL LOGOFF AT 14:46:58 ON 02 NOV 2006

Connection closed by remote host END

Unable to generate the STN prompt. Exiting the script...

Connecting via Winsock to STN

Welcome to STN International! Enter x:x

LOGINID: ssptalxn1621

PASSWORD:

TERMINAL (ENTER 1, 2, 3, OR ?):2

```
NEWS 1
                Web Page for STN Seminar Schedule - N. America
                CHEMLIST enhanced with New Zealand Inventory of Chemicals
NEWS 2 JAN 08
                CA/CAplus Company Name Thesaurus enhanced and reloaded
NEWS 3 JAN 16
                IPC version 2007.01 thesaurus available on STN
NEWS 4 JAN 16
NEWS 5 JAN 16 WPIDS/WPINDEX/WPIX enhanced with IPC 8 reclassification data
                CA/CAplus updated with revised CAS roles
NEWS 6 JAN 22
    7
                CA/CAplus enhanced with patent applications from India
        JAN 22
NEWS
NEWS 8 JAN 29
                PHAR reloaded with new search and display fields
                CAS Registry Number crossover limit increased to 300,000 in
NEWS 9 JAN 29
                multiple databases
NEWS 10 FEB 15
                PATDPASPC enhanced with Drug Approval numbers
NEWS 11 FEB 15 RUSSIAPAT enhanced with pre-1994 records
NEWS 12 FEB 23 KOREAPAT enhanced with IPC 8 features and functionality
NEWS 13 FEB 26 MEDLINE reloaded with enhancements
NEWS 14 FEB 26 EMBASE enhanced with Clinical Trial Number field
                TOXCENTER enhanced with reloaded MEDLINE
NEWS 15 FEB 26
       FEB 26
                IFICDB/IFIPAT/IFIUDB reloaded with enhancements
NEWS 16
                CAS Registry Number crossover limit increased from 10,000
NEWS 17 FEB 26
                 to 300,000 in multiple databases
                WPIDS/WPIX enhanced with new FRAGHITSTR display format
NEWS 18
        MAR 15
                CASREACT coverage extended
NEWS 19
        MAR 16
NEWS 20 MAR 20
                MARPAT now updated daily
NEWS 21 MAR 22 LWPI reloaded
                RDISCLOSURE reloaded with enhancements
NEWS 22 MAR 30
                JICST-EPLUS removed from database clusters and STN
NEWS 23 APR 02
NEWS 24 APR 30 GENBANK reloaded and enhanced with Genome Project ID field
NEWS 25
                CHEMCATS enhanced with 1.2 million new records
        APR 30
NEWS 26
                CA/CAplus enhanced with 1870-1889 U.S. patent records
        APR 30
                INPADOC replaced by INPADOCDB on STN
NEWS 27
        APR 30
NEWS 28 MAY 01
                New CAS web site launched
                CA/CAplus Indian patent publication number format defined
NEWS 29
        80 YAM
                RDISCLOSURE on STN Easy enhanced with new search and display
NEWS 30 MAY 14
                BIOSIS reloaded and enhanced with archival data
NEWS 31 MAY 21
```

NEWS 32 MAY 21 TOXCENTER enhanced with BIOSIS reload